=> d abs fbib 1-15

L7 ANSWER 1 OF 15 CAPLUS COPYRIGHT 2005 ACS on STN
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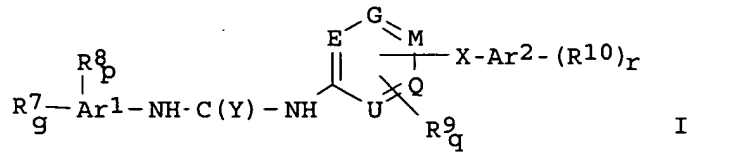

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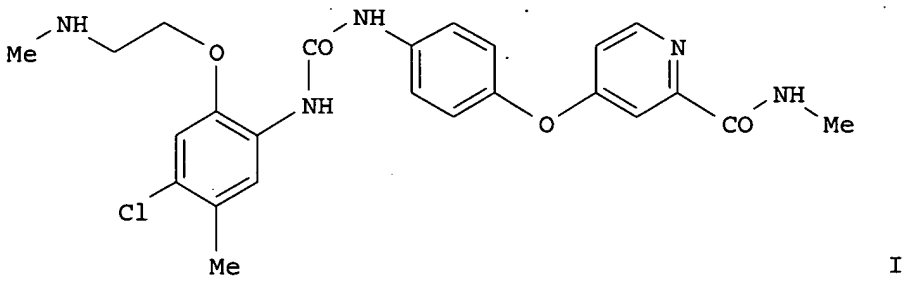

I

AB The present invention relates to bisarylurea derivs. (shown as I; variables defined below; e.g. 4-[4-[3-[4-chloro-5-methyl-2-(2-methylaminoethoxy)phenyl]ureido]phenoxy]pyridine-2-carboxylic acid methylamide (shown as II)), their use as inhibitors of raf-kinase (no data) and for the manufacture of a pharmaceutical composition and a method of treatment, comprising administering said pharmaceutical composition to a patient. Methods of preparation are claimed and >100 example prepns. are included. For example, 1-[2-[2-[(tert-butoxycarbonyl)(methyl)amino]ethoxy]-5-(trifluoromethyl)phenyl]-3-[4-[[2-(methylcarbamoyl)pyridin-4-yl]oxy]phenyl]urea was prepared (87 %) by reacting tert-Bu [2-[2-amino-4-(trifluoromethyl)phenoxy]ethyl](methyl)carbamate (preparation given) with p-nitrophenyl chloroformate followed by N-methyl-4-(4-aminophenoxy)pyridine-2-carboxamide (preparation given) and DIPEA; deprotection gave 86 % 1-[2-[2-(methylamino)ethoxy]-5-(trifluoromethyl)phenyl]-3-[4-[[2-(methylcarbamoyl)pyridin-4-yl]oxy]phenyl]urea. For I: Ar1, Ar2 = aromatic hydrocarbons containing 6 to 14 C atoms and ethylenic unsatd. or aromatic heterocyclic residues containing 3 to 10 C atoms and one or two heteroatoms, = N, O and S; E, G, M, Q and U = C and N atoms, with the proviso that ≥1 of E, G, M, Q and U are C atoms and that X is bonded to a C atom. R7 = Het, OHet, N(R11)Het, (CR5R6)kHet, et al. or R7 = -SO2-CR8:CR8-, wherein both valencies are bound vicinally to Ar1; R8, R9 and R10 = H, A, cycloalkyl comprising 3 to 7 C atoms, Hal, et al.; Y = O, S, NR21, C(R22)-NO2, C(R22)-CN and C(CN)2; g = 1-3, preferably 1 or 2, p, r = 0-5; q = 0-4, preferably 0, 1 or 2; addnl. details are given in the claims.

AN 2005:823661 CAPLUS
DN 143:229726

TI Preparation of 1,3-diarylureas as inhibitors of raf and other kinases useful against cancer and other diseases

IN Buchstaller, Hans-Peter; Burgdorf, Lars; Stieber, Frank; Amendt, Christiane; Grell, Matthias; Sirrenberg, Christian; Zenke, Frank

PA Merck Patent G.m.b.H., Germany
SO PCT Int. Appl., 264 pp.
CODEN: PIXXD2
DT Patent
LA English
FAN.CNT 1

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI WO 2005075425	A2	20050818	WO 2005-EP387	20050117

W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW
RW: BW, GH, GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IS, IT, LT, LU, MC, NL, PL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG

EP 2004-2092 A 20040130

L7 ANSWER 2 OF 15 CAPLUS COPYRIGHT 2005 ACS on STN
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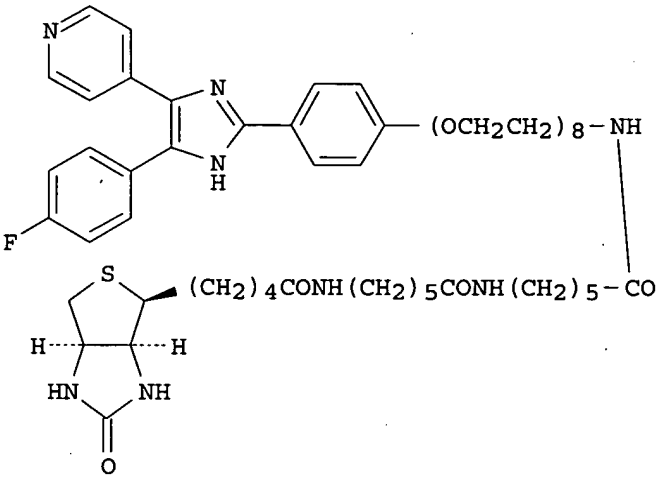

I

AB Provided herein are linker compds. and conjugates that include the linker compds. In one embodiment, the linker compds. comprise 2 or 3 residues of 6-aminohexanoic acid and optionally 7-10 residues of polyethyleneglycol (PEG). The linker compds. are useful in forming conjugates with one or more components useful in biopharmaceutical or bioanal. applications. In particular, the biopharmaceutically useful compds. are kinase inhibitors. The conjugates described herein have utility in a variety of diagnostic, separation, and therapeutic applications. Thus, I was prepared from SB 202190, PEG-azide and the biotin-linker compound
AN 2005:614536 CAPLUS
DN 143:115392

12/19/05

TI Preparation of conjugated small molecules for diagnostic and therapeutic use
IN Grotzfeld, Robert M.; Milanov, Zdravko V.; Patel, Hitesh K.; Lai, Andiliy G.; Mehta, Shamal A.; Lockhart, David J.
PA Ambit Biosciences Corp., USA
SO U.S. Pat. Appl. Publ., 63 pp.
CODEN: USXXCO
DT Patent
LA English
FAN.CNT 1

PATENT NO.	KIND	DATE	APPLICATION NO.		DATE
PI US 2005153371	A1	20050714	US 2005-31638		20050107
			US 2004-535173P	P	20040107
			US 2004-557941P	P	20040330
WO 2005067644	A2	20050728	WO 2005-US456		20050107
WO 2005067644	A3	20051013			

W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW
RW: BW, GH, GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IS, IT, LT, LU, MC, NL, PL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG

US 2004-535173P	P	20040107		
US 2004-557941P	P	20040330		

L7 ANSWER 3 OF 15 CAPLUS COPYRIGHT 2005 ACS on STN
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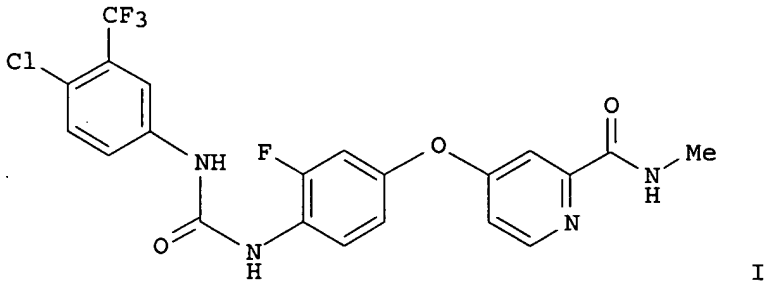

I

AB Title compound I is prepared I and salts thereof is prepared in several steps from 3-fluoro-4-nitrophenol, 4-chloro-N-methylpyridine-2-carboxamide and 4-chloro-3-(trifluoromethyl)phenylisocyanate. I inhibits PDGFR tyrosine kinase with IC50 = 83nM. I is useful for the treatment of, e.g., inflammation and as an antiproliferative agent.
AN 2005:99470 CAPLUS
DN 142:197889
TI Fluoro substituted omega-carboxyaryl diphenyl urea for treatment of raf, VEGFR, PDGFR, p38 and flt-3 kinase-mediated diseases
IN Dumas, Jacques; Boyer, Stephen; Riedl, Bernd; Wilhelm, Scott
PA Bayer Pharmaceuticals Corporation, USA

SO PCT Int. Appl., 68 pp.
CODEN: PIXXD2
DT Patent
LA English
FAN.CNT 1

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI WO 2005009961	A2	20050203	WO 2004-US23500	20040722
WO 2005009961	A3	20050331		
WO 2005009961	B1	20050602		

W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW
RW: BW, GH, GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG

			US 2003-489102P	P 20030723
			US 2004-540326P	P 20040202
US 2005038080	A1	20050217	US 2004-895985	20040722
			US 2003-489102P	P 20030723
			US 2004-540326P	P 20040202

L7 ANSWER 4 OF 15 CAPLUS COPYRIGHT 2005 ACS on STN
AB The present invention provides methods for treating and/or preventing conditions and diseases in humans and other mammals that are associated with and/or mediated by signal transduction pathways comprising platelet-derived growth factor receptor (PDGFR), especially PDGFR-β, by administering diaryl ureas. The present invention also provides devices and methods for treating, ameliorating, preventing, or modulating restenosis following angioplastic surgery or other invasive procedures that affect or injure the vascular system, and graft rejection following transplantation of a donor tissue into a host, where a stent or other implantable device comprises an effective amount of diaryl ureas. For example, N-[4-chloro-3-(trifluoromethyl)phenyl]-N'-[4-[2-(N-methylcarbamoyl)-4-pyridyloxy]phenyl] urea, N-[4-chloro-3-(trifluoromethyl)phenyl]-N'-[4-[2-(N-methylcarbamoyl)-4-pyridyloxy]-2-fluorophenyl] urea, and N-[4-chloro-3-(trifluoromethyl)phenyl]-N'-[4-[2-(N-methylcarbamoyl)-4-pyridyloxy]-2-chl orophenyl]urea showed an IC50 of less than 10 μM in a pPDGFR-β sandwich ELISA in AoSMC cells.
AN 2005:14200 CAPLUS
DN 142:86701
TI Diaryl ureas for treatment of diseases mediated by PDGFR
IN Wilhelm, Scott; Dumas, Jacques; Ladouceur, Gaetan; Lynch, Mark; Scott, William J.
PA Bayer Pharmaceuticals Corporation, USA
SO PCT Int. Appl., 47 pp.
CODEN: PIXXD2
DT Patent
LA English
FAN.CNT 2

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI WO 2005000284	A2	20050106	WO 2004-US15653	20040519
WO 2005000284	A3	20050310		

W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW
RW: BW, GH, GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG

				US 2003-471735P	P 20030520
				US 2003-520399P	P 20031117
				US 2004-556062P	P 20040325
US 2005059703	A1	20050317		US 2004-848567	20040519
				US 2003-471735P	P 20030520
				US 2003-520399P	P 20031117
				US 2004-556062P	P 20040325

PATENT FAMILY INFORMATION:

FAN 2004:1154653

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI WO 2004113274	A2	20041229	WO 2004-US15655	20040519
WO 2004113274	A3	20050303		

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RW: BW, GH, GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG

				US 2003-471735P	P 20030520
				US 2003-520399P	P 20031117
				US 2004-556062P	P 20040325
US 2005059703	A1	20050317		US 2004-848567	20040519
				US 2003-471735P	P 20030520
				US 2003-520399P	P 20031117
				US 2004-556062P	P 20040325

OS MARPAT 142:86701

L7 ANSWER 5 OF 15 CAPLUS COPYRIGHT 2005 ACS on STN

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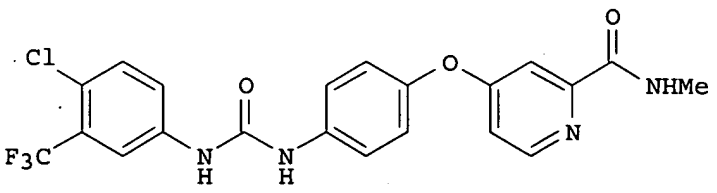

I

AB Diaryl ureas B-NH-CO-NH-L-(CH2)m-X-(CH2)p-L1-(Q)1-3 [I; B = (un)substituted Ph, naphthyl, or heteroaryl; L, =(un)substituted Ph, naphthyl, or heteroaryl; X = bond, O, CO, NR3, NR3CO, S, CONR3, CF2, CCl2, CHF, CH(OH), C.tplbond.C, CH:CH, CR4R5; m, p = independently 0-4; L1 = any group L, 5-6 membered cyclic structure; Q = independently COR4, CO2R4, CONR4R5; each R3-R5 = independently H, (un)substituted C1-5 alkyl,C3-5 cycloalkyl, Ph, C1-3 alkylphenyl, C0-4 alkylheteroaryl], useful to treat diseases and conditions associated with signal transduction pathways comprising of at least one of raf, VEGFR, PDGFR, p38 and/or FLT-3. E.g., a multi-step synthesis of the urea II which produced dose-dependent 45-68% inhibition of tumor growth in a staged HCT 116 colon (mutant k-Ras) xenograft model.

AN 2004:1154653 CAPLUS

DN 142:93545

TI Preparation of diaryl ureas with kinase inhibiting activity

IN Wilhelm, Scott; Dumas, Jacques; Ladouceur, Gaetan; Lynch, Mark; Scott, William J.

PA Bayer Pharmaceuticals Corporation, USA

SO PCT Int. Appl., 122 pp.
CODEN: PIXXD2

DT Patent

LA English

FAN.CNT 2

	PATENT NO.	KIND	DATE	APPLICATION NO.		DATE
PI	WO 2004113274	A2	20041229	WO 2004-US15655		20040519
	WO 2004113274	A3	20050303			
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	RW: BW, GH, GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG					
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				US 2003-520399P	P	20031117
				US 2004-556062P	P	20040325
	US 2005059703	A1	20050317	US 2004-848567		20040519
				US 2003-471735P	P	20030520
				US 2003-520399P	P	20031117
				US 2004-556062P	P	20040325

PATENT FAMILY INFORMATION:

FAN 2005:14200

	PATENT NO.	KIND	DATE	APPLICATION NO.		DATE
PI	WO 2005000284	A2	20050106	WO 2004-US15653		20040519
	WO 2005000284	A3	20050310			
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AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG

			US 2003-471735P	P	20030520
			US 2003-520399P	P	20031117
			US 2004-556062P	P	20040325
US 2005059703	A1	20050317	US 2004-848567		20040519
			US 2003-471735P	P	20030520
			US 2003-520399P	P	20031117
			US 2004-556062P	P	20040325

OS MARPAT 142:93545

L7 ANSWER 6 OF 15 CAPLUS COPYRIGHT 2005 ACS on STN

AB Three-dimensional quant. structure activity relationship (3D-QSAR) analyses were carried out on 91 substituted ureas in order to understand their Raf-1 kinase inhibitory activities. The studies include Comparative Mol. Field Anal. (Co-MFA) and Comparative Mol. Similarity Indexes Anal. (Co-MSIA). Models with good predictive abilities were generated with the cross validated r2 (r2cv) values for Co-MFA and Co-MSIA being 0.53 and 0.44, resp. The conventional r2 values are 0.93 and 0.87 for Co-MFA and Co-MSIA, resp. In addition, a homol. model of Raf-1 was also constructed using the crystal structure of the kinase domain of B-Raf isoform with one of the most active Raf-1 inhibitors (I) inside the active site. The ATP binding pocket of Raf-1 is virtually similar to that of B-Raf. Selected ligands were docked in the active site of Raf-1. Mol. I adopts an orientation similar to that inside the B-Raf active site. The 4-pyridyl group bearing amide substituent is located in the adenosine binding pocket, and anchored to the protein through a pair of hydrogen bonds with Cys424 involving ring N-atom and amide NH group. The results of best 3D-QSAR model were compared with structure-based studies using the Raf-1 homol. model. The results of 3D-QSAR and docking studies validate each other and provided insight into the structural requirements for activity of this class of mols. as Raf-1 inhibitors. Based on these results, novel mols. with improved activity can be designed.

AN 2004:1004044 CAPLUS

DN 142:126552

TI 3D-QSAR CoMFA, CoMSIA studies on substituted ureas as Raf-1 kinase inhibitors and its confirmation with structure-based studies

AU Thaimattam, Ram; Daga, Pankaj; Rajjak, Shaikh Abdul; Banerjee, Rahul; Iqbal, Javed

CS Department of Molecular Modeling and Drug Design, Discovery Research, Dr. Reddy's Laboratories Ltd., Hyderabad, 500 049, India

SO Bioorganic & Medicinal Chemistry (2004), 12(24), 6415-6425
CODEN: BMECEP; ISSN: 0968-0896

PB Elsevier Ltd.

DT Journal

LA English

RE.CNT 29 THERE ARE 29 CITED REFERENCES AVAILABLE FOR THIS RECORD
ALL CITATIONS AVAILABLE IN THE RE FORMAT

L7 ANSWER 7 OF 15 CAPLUS COPYRIGHT 2005 ACS on STN

AB Urea derivs. of formula A-NHCONH-B or pharmaceutically acceptable salts thereof [A = a substituted moiety of up to 40 carbon atoms of the formula -L-(M-L1)q; where L = a 5 or 6 membered cyclic structure bound directly to D; L1 = a substituted cyclic moiety having at least 5 members; M = a bridging group having at least one atom; q = an integer of 1-3; each cyclic structure of L and L1 contains 0-4 members of the group consisting of nitrogen, oxygen and sulfur; B = a substituted or unsubstituted, up to

tricyclic aryl or heteroaryl moiety of up to 30 carbon atoms with at least one 6-member cyclic structure bound directly to D containing 0-4 members of the group consisting of nitrogen, oxygen and sulfur] are prepared These compds. are useful for raf mediated diseases, in particular a cancerous cell growth mediated by raf kinase. All compds. exemplified, e.g. N-[4-Chloro-3-(trifluoromethyl)phenyl]-N'-[4-[2-(N-methylcarbamoyl)-4-pyridyloxy]phenyl]urea, displayed IC50 of between 1 mM and 10 μM.

AN 2003:874965 CAPLUS
DN 139:364958
TI Preparation of omega-carboxyaryl substituted diphenyl ureas as raf kinase inhibitors
IN Riedl, Bernd; Dumas, Jacques; Khire, Uday; Lowinger, Timothy B.; Scott, William J.; Smith, Roger A.; Wood, Jill E.; Monahan, Mary-Katherine; Natero, Reina; Renick, Joel; Sibley, Robert N.
PA Bayer Corporation, USA
SO U.S. Pat. Appl. Publ., 60 pp.
CODEN: USXXCO
DT Patent
LA English
FAN.CNT 1

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI US 2003207872	A1	20031106	US 2002-42226	20020111
			US 2002-42226	20020111

OS MARPAT 139:364958

L7 ANSWER 8 OF 15 CAPLUS COPYRIGHT 2005 ACS on STN
AB Aryl ureas of formula A-NHCONH-B [A = a substituted moiety of up to 40 carbon atoms of the formula: -L-(M-L1)q (where L = a 5 or 6 membered cyclic structure bound directly to D, L1 comprises a substituted cyclic moiety having at least 5 members; M = a bridging group having at least one atom; q = an integer of from 1-3; each cyclic structure of L and L1 contains 0-4 members of the group consisting of nitrogen, oxygen and sulfur); B = a substituted or unsubstituted, up to tricyclic aryl or heteroaryl moiety of up to 30 carbon atoms with at least one 6-member cyclic structure bound directly to D containing 0-4 members of the group consisting of nitrogen, oxygen and sulfur] are prepared These urea derivs. are useful for treating raf mediated diseases, in particular cancerous cell growth mediated by raf kinase. Thus, N-[4-bromo-3-(trifluoromethyl)phenyl]-N'-[4-[2-(N-methylcarbamoyl)-4-pyridyloxy]phenyl]urea. Thus, a solution of 4-bromo-3-(trifluoromethyl)phenyl isocyanate (8.0 g, 30.1 mmol) in CH2Cl2 (80 mL) was added dropwise to a solution of 4-[2-(N-methylcarbamoyl)-4-pyridyloxy]aniline (7.0 g, 28.8 mmol) in CH2Cl2 (40 mL) at 0°, stirred at room temperature for 16 h, and filtered to give, after washing the yellow solids, washing with CH2Cl2 (2 + 50 mL), and drying under reduced pressure (approx. 1 mmHg) at 40° to give N-[4-bromo-3-(trifluoromethyl)phenyl]-N'-[4-[2-(N-methylcarbamoyl)-4-pyridyloxy]phenyl]urea. All compds. exemplified showed IC50 between 1 nM to 10 μM against raf kinase.

AN 2003:757329 CAPLUS
DN 139:276918
TI Preparation of omega-carboxyaryl substituted diphenyl ureas as raf kinase inhibitors
IN Riedl, Bernd; Dumas, Jacques; Khire, Uday; Lowinger, Timothy B.; Scott, William J.; Smith, Roger A.; Wood, Jill E.; Monahan, Mary-katherine; Natero, Reina; Renick, Joel; Sibley, Robert N.
PA Bayer Corporation, USA
SO U.S. Pat. Appl. Publ., 61 pp.

CODEN: USXXCO
DT Patent
LA English
FAN.CNT 1

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI US 2003181442	A1	20030925	US 2001-993647	20011127
			US 2001-993647	20011127

OS MARPAT 139:276918

L7 ANSWER 9 OF 15 CAPLUS COPYRIGHT 2005 ACS on STN
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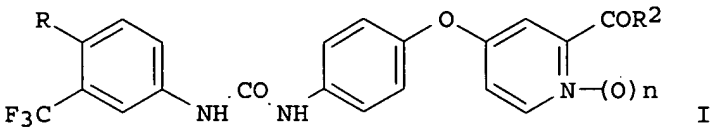

I

AB Aryl ureas, such as I [R = Cl, Br; R2 = OH, NH2, NHMe, NHCH2OH, alkoxy; n = 0, 1], were prepared for use in pharmaceutical compns. for the treatment of raf kinase and p38 kinase mediated diseases. These ureas are useful for the treatment of inflammation, osteoporosis, angiogenesis disorders and hyper-proliferative disorders, such as cancer. Thus, urea I (R = Cl, R2 = NHMe, n = 1) was prepared with 57% yield by N-oxidation of I (R = Cl, R2 = NHMe, n = 0) using 3-chloroperbenzoic acid in CH2Cl2 and THF. The prepared ureas were assayed for inhibition of p38 kinase and raf kinase, as well as for cancer cell growth inhibition in human cancer cell lines, such as HCT116 and DLD-1.

AN 2003:656745 CAPLUS
DN 139:197377
TI Preparation of aryl ureas for therapeutic use as kinase inhibitors
IN Dumas, Jacques; Scott, William J.; Chien, Du-Schieng; Lee, Wendy; Bjorge, Susan; Musza, Laszlo L.; Nassar, Ala; Riedl, Bernd
PA Bayer Corporation, USA; Bayer Pharmaceuticals Corporation
SO PCT Int. Appl., 64 pp.
CODEN: PIXXD2
DT Patent
LA English
FAN.CNT 1

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI WO 2003068746	A1	20030821	WO 2003-US4109	20030211
WO 2003068746	C1	20050506		
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RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PT, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG				
			US 2002-354937P	P 20020211
CA 2475818	AA	20030821	CA 2003-2475818	20030211

			US 2002-354937P	P 20020211
			WO 2003-US4109	W 20030211
US 2003216446	A1	20031120	US 2003-361859	20030211
			US 2002-354937P	P 20020211
EP 1474393	A1	20041110	EP 2003-707848	20030211
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			US 2002-354937P	P 20020211
			WO 2003-US4109	W 20030211
JP 2005523278	T2	20050804	JP 2003-567877	20030211
			US 2002-354937P	P 20020211
			WO 2003-US4109	W 20030211
EP 1580188	A1	20050928	EP 2005-7027	20030211
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			US 2002-354937P	P 20020211
			EP 2003-707848	A3 20030211

OS MARPAT 139:197377
RE.CNT 2 THERE ARE 2 CITED REFERENCES AVAILABLE FOR THIS RECORD
ALL CITATIONS AVAILABLE IN THE RE FORMAT

L7 ANSWER 10 OF 15 CAPLUS COPYRIGHT 2005 ACS on STN
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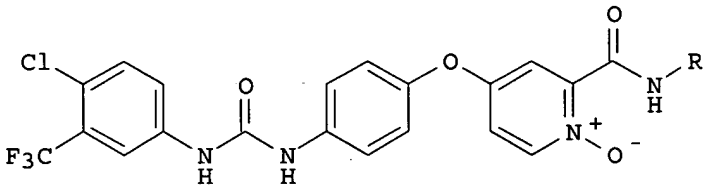

I

AB The title ureas containing a pyridine, quinoline, or isoquinoline functionality which is oxidized at the nitrogen heteroatom MLBNHCONHA [A = (un)substituted Ph, naphthyl, 5-6 membered monocyclic heteroaryl, 8-10 membered bicyclic heteroaryl; B = (un)substituted phenylene, naphthylene, 5-6 membered monocyclic heteroarylene, 8-10 membered bicyclic heteroarylene; L = (CH2)mO(CH2)l, (CH2)m(CH2)l, (CH2)mCO(CH2)l, etc.; m, l = 0-4; M = (un)substituted pyridine-1-oxide, quinoline-1-oxide, isoquinoline-1-oxide; with the provisos] which are useful in the treatment of (i) raf mediated diseases, for example, cancer, (ii) p38 mediated diseases such as inflammation and osteoporosis, and (iii) VEGF mediated diseases such as angiogenesis disorders, were claimed. Preparation of two ureas such as I [R = H, Me] which are not compds. of the invention, and have been distinguished from the compds. of the invention by a proviso, was described. Pharmaceutical composition comprising the title ureas was claimed.

AN 2003:656581 CAPLUS
DN 139:197370
TI Preparation of aryl ureas containing pyridine, quinoline and isoquinoline N-oxide functionality as kinase inhibitors
IN Dumas, Jacques; Scott, William J.; Riedl, Bernd
PA Bayer Corporation, USA
SO PCT Int. Appl., 67 pp.
CODEN: PIXXD2
DT Patent

LA English
FAN.CNT 1

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI WO 2003068229	A1	20030821	WO 2003-US4110	20030211
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RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PT, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG				
			US 2002-354935P	P 20020211
US 2003216396	A1	20031120	US 2003-361850	20030211
			US 2002-354935P	P 20020211

OS MARPAT 139:197370
RE.CNT 2 THERE ARE 2 CITED REFERENCES AVAILABLE FOR THIS RECORD
ALL CITATIONS AVAILABLE IN THE RE FORMAT

L7 ANSWER 11 OF 15 CAPLUS COPYRIGHT 2005 ACS on STN
AB ADB [I; D = NHCONH; A = L(ML1)q; L = 5-6 membered cyclic structure bound directly to D; L1 = substituted cyclic moiety having ≥5 members, M = bridging group having ≥1 atom; q = 1-3; L, L1 contain 0-4 N, O, S; B = (substituted) up to tricyclic aryl, heteroaryl of ≤30 C atoms with ≥1 6-membered cyclic structure bound directly to D containing 0-4 N, O, S], were prepared Thus, 4-chloro-3-(trifluoromethyl)phenyl isocyanate in CH2Cl2 was added dropwise to a suspension of 4-[2-(N-methylcarbamoyl)-4-pyridyloxy]aniline (preparation given) in CH2Cl2 at 0°; the resulting mixture was stirred at room temperature for 22 h. to afford N-[4-chloro-3-(trifluoromethyl)phenyl]-N'-[4-[2-(N-methylcarbamoyl)-4-pyridyloxy]phenyl]urea. I inhibited RAF kinase in the range 1 nM-1 μM. I pharmaceutical compns. are claimed.
AN 2003:590832 CAPLUS
DN 139:149528
TI Preparation of diphenylureas as RAF kinase inhibitors
IN Riedl, Bernd; Dumas, Jacques; Khire, Uday; Lowinger, Timothy B.; Scott, William J.; Smith, Roger A.; Wood, Jill E.; Monahan, Mary-katherine; Natero, Reina; Renick, Joel; Sibley, Robert N.
PA Bayer Corporation, USA
SO U.S. Pat. Appl. Publ., 62 pp., Cont. of U. S. Ser. No. 42,203.
CODEN: USXXCO
DT Patent
LA English
FAN.CNT 1

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI US 2003144278	A1	20030731	US 2002-283248	20021030
			US 2001-367380P	P 20010112
			US 2002-42203	A1 20020111

OS MARPAT 139:149528

L7 ANSWER 12 OF 15 CAPLUS COPYRIGHT 2005 ACS on STN
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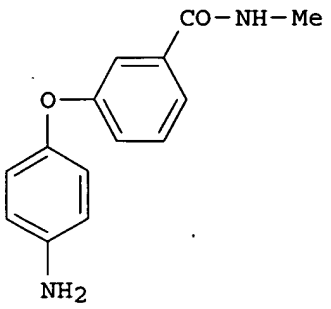

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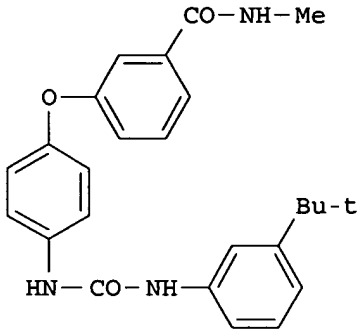

III

AB Title compds. B-NHCONH-L-(M-L1)q (I) [B = (un)substituted pyridyl, quinolinyl, isoquinolinyl; L = 5 or 6 membered cyclic structure; L1 = substituted cyclic moiety having at least 5 members; M = bridging group having at least one atom; q = 1-3; with proviso that L and L1 contain 0-4 hetero atoms, e.g., N, O and S] and their pharmaceutically acceptable salts were prepared For example, coupling of aniline II, e.g., prepared from Et 3-hydroxybenzoate in 4-steps, with bis(trichloromethyl)carbonate followed by 3-tert-butylaniline afforded urea III. In in vitro raf kinase assays, 112-specific examples of compds. I inhibited kinase activity with IC50 values ranging from 10 nM-10 μM. Compds. I are useful for the treatment of cancerous cell growth mediated by raf kinase.

AN 2002:850357 CAPLUS

DN 137:352907

TI Preparation of quinolyl, isoquinolyl or pyridyl-ureas as inhibitors of raf kinase for the treatment of tumors and/or cancerous cell growth

IN Dumas, Jacques; Riedl, Bernd; Khire, Uday; Wood, Jill E.; Robert, Sibley N.; Monahan, Mary-Katherine; Renick, Joel; Gunn, David E.; Lowinger, Timothy B.; Scott, William J.; Smith, Roger A.

PA Bayer Corporation, USA

SO U.S. Pat. Appl. Publ., 63 pp., Cont.-in-part of U.S. Ser. No. 758,548.
CODEN: USXXCO

DT Patent

LA English

FAN.CNT 5

	PATENT NO.	KIND	DATE	APPLICATION NO.		DATE
PI	US 2002165394	A1	20021107	US 2001-777920		20010207
				US 1999-115877P	P	19990113
				US 1999-257266	B2	19990225
				US 1999-425228	B2	19991022
				US 2001-758548	A2	20010112
	ZA 2001005751	A	20030714	ZA 2001-5751		20010712
				US 1999-115877P	P	19990113
	US 2002137774	A1	20020926	US 2001-907970		20010719
				US 1999-115877P	P	19990113
	WO 2002062763	A2	20020815	WO 2002-US3361		20020207
	WO 2002062763	A3	20021010			

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RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG

				US 2001-777920	A 20010207
US 2003139605	A1	20030724		US 2002-71248	20020211
				US 1999-115877P	P 19990113
				US 1999-115878P	P 19990113
				US 1999-257266	B2 19990225
				US 1999-425228	B1 19991022
				US 2001-948915	A1 20010910

PATENT FAMILY INFORMATION:

FAN 2000:493376

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI	WO 2000041698	A1	20000720	WO 2000-US768	20000113

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RW: GH, GM, KE, LS, MW, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG

				US 1999-115878P	P 19990113
				US 1999-257265	A2 19990225
				US 1999-425229	A2 19991022
CA 2359244	AA	20000720		CA 2000-2359244	20000113
				US 1999-115878P	P 19990113
				US 1999-257265	A 19990225
				US 1999-425229	A 19991022
				WO 2000-US768	W 20000113
EP 1158985	A1	20011205		EP 2000-905597	20000113

R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO

				US 1999-115878P	P 19990113
				US 1999-257265	A 19990225
				US 1999-425229	A 19991022
				WO 2000-US768	W 20000113
US 2003139605	A1	20030724		US 2002-71248	20020211
				US 1999-115877P	P 19990113
				US 1999-115878P	P 19990113
				US 1999-257266	B2 19990225
				US 1999-425228	B1 19991022
				US 2001-948915	A1 20010910
US 2003105091	A1	20030605		US 2002-86417	20020304
				US 1999-115878P	P 19990113
				US 1999-257265	B2 19990225
				US 1999-425229	B1 19991022

FAN 2000:493516

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI	WO 2000042012	A1	20000720	WO 2000-US648	20000112

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			US 1999-257266	A2	19990225
			US 1999-425228	A2	19991022
CA 2359510	AA	20000720	CA 2000-2359510		20000112
			US 1999-115877P	P	19990113
			US 1999-257266	A	19990225
			US 1999-425228	A	19991022
			WO 2000-US648	W	20000112
AU 2000025016	A5	20000801	AU 2000-25016		20000112
			US 1999-115877P	P	19990113
			US 1999-257266	A	19990225
			US 1999-425228	A	19991022
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EP 1140840	A1	20011010	EP 2000-903239		20000112

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			US 1999-257266	A	19990225
			US 1999-425228	A	19991022
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			US 1999-257266	A	19990225
			US 1999-425228	A	19991022
			WO 2000-US648	W	20000112
JP 2003526613	T2	20030909	JP 2000-593580		20000112
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			US 1999-425228	A	19991022
			WO 2000-US648	W	20000112
BR 2000007487	A	20030923	BR 2000-7487		20000112
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			US 1999-257266	A	19990225
			US 1999-425228	A	19991022
			WO 2000-US648	W	20000112
US 2001011135	A1	20010802	US 2001-773659		20010202
			US 1999-115877P	P	19990113
			US 1999-257266	B2	19990225
			US 1999-425228	A1	19991022
US 2001011136	A1	20010802	US 2001-773675		20010202
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			US 1999-257266	B2	19990225
			US 1999-425228	A1	19991022
US 2001016659	A1	20010823	US 2001-773672		20010202
			US 1999-115877P	P	19990113
			US 1999-257266	B2	19990225
			US 1999-425228	A1	19991022
US 2001027202	A1	20011004	US 2001-773658		20010202
			US 1999-115877P	P	19990113
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			US 1999-115877P	P	19990113
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				US 1999-257266	A	19990225
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	US 2002137774	A1	20020926	US 2001-907970		20010719
				US 1999-115877P	P	19990113
	BG 105763	A	20020329	BG 2001-105763		20010801
				US 1999-115877P	P	19990113
				US 1999-257266	A	19990225
				US 1999-425228	A	19991022
				WO 2000-US648	W	20000112
	HR 2001000580	A1	20020831	HR 2001-580		20010802
				US 1999-115877P	P	19990113
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	US 2002042517	A1	20020411	US 2001-948915		20010910
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				US 1999-257266	B2	19990225
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	US 2003139605	A1	20030724	US 2002-71248		20020211
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				US 1999-115878P	P	19990113
				US 1999-257266	B2	19990225
				US 1999-425228	B1	19991022
				US 2001-948915	A1	20010910

FAN 2002:409267

	PATENT NO.	KIND	DATE	APPLICATION NO.		DATE
PI	US 2002065296	A1	20020530	US 2001-838286		20010420
				US 1999-115878P	P	19990113
				US 1999-257265	B1	19990225
				US 1999-425229	A2	19991022
				US 2001-778039	A2	20010207
	US 2003139605	A1	20030724	US 2002-71248		20020211
				US 1999-115877P	P	19990113
				US 1999-115878P	P	19990113
				US 1999-257266	B2	19990225
				US 1999-425228	B1	19991022
				US 2001-948915	A1	20010910
	CA 2443952	AA	20021031	CA 2002-2443952		20020417
				US 2001-838286	A	20010420
				WO 2002-US12064	W	20020417
	WO 2002085859	A1	20021031	WO 2002-US12064		20020417
	W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PH, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM					
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				US 2001-838286	A	20010420
	EP 1379507	A1	20040114	EP 2002-725709		20020417
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				US 2001-838286	A	20010420
				WO 2002-US12064	W	20020417
	JP 2004537511	T2	20041216	JP 2002-583386		20020417
				US 2001-838286	A	20010420
				WO 2002-US12064	W	20020417

FAN 2002:615574

	PATENT NO.	KIND	DATE	APPLICATION NO.		DATE
PI	WO 2002062763	A2	20020815	WO 2002-US3361		20020207
	WO 2002062763	A3	20021010			

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	US 2002165394	A1	20021107	US 2001-777920		20010207
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				US 1999-257266	B2	19990225
				US 1999-425228	B2	19991022
				US 2001-758548	A2	20010112

OS MARPAT 137:352907

L7 ANSWER 13 OF 15 CAPLUS COPYRIGHT 2005 ACS on STN

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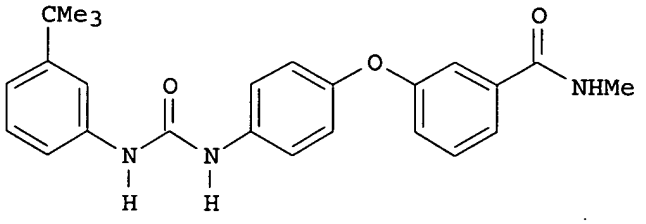

II

AB Title compds., e.g., RNHCONHZOR1 [I; R = C6H4(CMe3)-3, 2-methoxy-5-trifluoromethylphenyl, 4-chloro-3-trifluoromethylphenyl, 2-methoxy-3-quinolyl, etc.; R1 = (un)substituted acylphenyl, -acylpyridinyl, etc.; Z = (un)substituted 1,3- or -1,4-phenylene] were prepared Thus, 4-(H2N)C6H4OC6H4(CONHMe)-4 (preparation given) was condensed with 3-(Me3C)C6H4NH2 and CO(OCCl3)2 to give title compound II. Data for biol. activity of title compds. were given.

AN 2002:615574 CAPLUS

DN 137:169425

TI Preparation of N-aryl-N'-[(acylphenoxy)phenyl]ureas as raf kinase inhibitors

IN Dumas, Jacques; Riedl, Bernd; Khire, Uday; Wood, Jill E.; Sibley, Robert N.; Monahan, Mary-Katherine; Renick, Joel; Gunn, David E.; Lowinger, Timothy B.; Scott, William J.; Smith, Roger A.

PA Bayer Corporation, USA
SO PCT Int. Appl., 125 pp.
CODEN: PIXXD2
DT Patent
LA English
FAN.CNT 5

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
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				US 1999-425228	B2 19991022
				US 2001-758548	A2 20010112

PATENT FAMILY INFORMATION:

FAN 2000:493376

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FAN 2000:493516

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FAN 2002:409267

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FAN 2002:850357

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OS MARPAT 137:169425

L7 ANSWER 14 OF 15 CAPLUS COPYRIGHT 2005 ACS on STN
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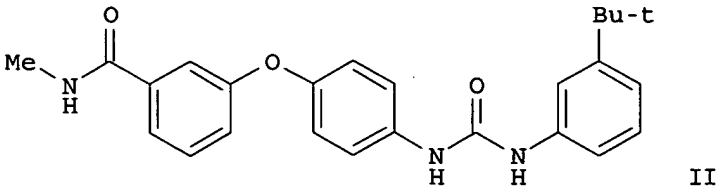

II

AB This invention relates to the preparation and use of (hetero)aryl ureas ANHCONHB [I; A = L(ML1)q; L = 5- or 6-membered (hetero)aryl, especially Ph or pyridinyl; M = bridging group; L1 = (hetero)aryl with at least one (un)substituted sulfamoyl, carboxy, or carbamoyl substituent; q = 1-3; B = certain (un)substituted mono- to tricyclic aryl or heteroaryl groups] for the treatment of raf mediated diseases, such as cancer (no data). Approx. 100 invention compds. and numerous intermediates were prepared For instance, 3-tert-butylaniline was coupled with bis(trichloromethyl)carbonate to form the isocyanate, followed by addition of 4-(3-N-methylcarbamoylphenoxy)aniline (preparation given) to afford the urea II.

AN 2000:493516 CAPLUS

DN 133:120157

TI Preparation of ω-carboxy(hetero)aryl substituted diphenyl ureas as raf kinase inhibitors

IN Riedl, Bernd; Dumas, Jacques; Khire, Uday; Lowinger, Timothy B.; Scott, William J.; Smith, Roger A.; Wood, Jill E.; Monahan, Mary-Katherine; Natero, Reina; Renick, Joel; Sibley, Robert N.

PA Bayer Corporation, USA

SO PCT Int. Appl., 120 pp.
CODEN: PIXXD2

DT Patent

LA English

FAN.CNT 5

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PATENT FAMILY INFORMATION:

FAN 2000:493376

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				US 1999-257265	A2	19990225
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				US 1999-425228	B1	19991022
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	US 2003105091	A1	20030605	US 2002-86417		20020304
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FAN 2002:409267

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				US 1999-115878P	P	19990113
				US 1999-257266	B2	19990225
				US 1999-425228	B1	19991022
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				WO 2002-US12064	W	20020417
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FAN 2002:615574

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US 2003139605	A1	20030724		US 2002-71248	20020211
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				US 1999-115878P	P 19990113
				US 1999-257266	B2 19990225
				US 1999-425228	B1 19991022
				US 2001-948915	A1 20010910

OS MARPAT 133:120157
RE.CNT 12 THERE ARE 12 CITED REFERENCES AVAILABLE FOR THIS RECORD
ALL CITATIONS AVAILABLE IN THE RE FORMAT

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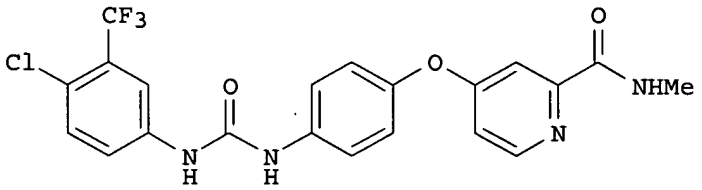

II

AB The title compds. ADB [I; D = NHCONH; A = substituted moiety of up to 40 carbon atoms of the formula L(ML1)q (wherein L = 5-6 membered cyclic structure; L1 = substituted cyclic moiety having at least 5 members; M = bridging group having al least one atom; q = 1-3; each of L and L1 contains 0-4 members of the group consisting of N, O and S); B = (un)substituted up to tricyclic aryl or heteroaryl moiety of up to 30 carbon atoms with at least one 6-member cyclic structure bound directly to D containing 0-4 members of the group consisting of N, O and S], useful in treating p38 mediated diseases, were prepared E.g., a multi-step synthesis of the urea II which showed IC50 of 1-10 μM against p38, was given. Compds. I are effective at 0.01-200 mg/kg/day (oral administration).

AN 2000:493376 CAPLUS
DN 133:120155
TI Preparation of ω-carboxy aryl substituted diphenyl ureas as p38 kinase inhibitors
IN Riedl, Bernd; Dumas, Jacques; Khire, Uday; Lowinger, Timothy B.; Scott, William J.; Smith, Roger A.; Wood, Jill E.; Monahan, Mary-Katherine; Natero, Reina; Renick, Joel; Sibley, Robert N.
PA Bayer Corporation, USA
SO PCT Int. Appl., 148 pp.
CODEN: PIXXD2
DT Patent

LA English
FAN.CNT 5

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PI	WO 2000041698	A1	20000720	WO 2000-US768		20000113
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				US 1999-425228	B1	19991022
				US 2001-948915	A1	20010910
	US 2003105091	A1	20030605	US 2002-66417		20020304
				US 1999-115878P	P	19990113
				US 1999-257265	B2	19990225
				US 1999-425229	B1	19991022

PATENT FAMILY INFORMATION:
FAN 2000:493516

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				WO 2000-US648	W	20000112
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FAN 2002:409267

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RE.CNT 1 THERE ARE 1 CITED REFERENCES AVAILABLE FOR THIS RECORD
ALL CITATIONS AVAILABLE IN THE RE FORMAT